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☐ 1: Ann N Y Acad Sci. 1998 Jun 30;850:163-77.



Full Text
Ann N Y Acad Sci

Link

Targeted integration of a recombinant globin gene adeno-associated viral vector into human chromosome 19.

Bertran J, Yang Y, Hargrove P, Vanin EF, Nienhuis AW.

Department of Hematology/Oncology, St. Jude Children's Research Hospital, Memphis, Tennessee 38105, USA.

Transfer of a globin gene into stem cells along with the regulatory elements required to achieve high level expression in maturing erythroid cells would provide effective gene therapy for Cooley's Anemia. We have explored the use of recombinant adeno-associated viral (rAAV) vectors for this purpose. A vector designated rHS32A gamma*3'RE that contains regulatory elements from the locus control and flanking regions, integrates as a stable head-to-tail concatamer in erythroleukemia cells at a high multiplicity of infection and exhibits high level, regulated gamma globin gene expression. Inducible expression of the non-structural Rep proteins of wild-type AAV in HeLa cells transduced with rAAV vectors does not increase overall integration frequency, but targeted integration of rHS32A gamma*3'RE into human chromosome 19 was documented.

PMID: 9668538 [PubMed - indexed for MEDLINE]

Related Links

High-level globin gene expression mediated by a recombinant adeno-associated virus genome that contains the 3' gamma globin gene regulatory element and integrates as tandem copies in erythroid cells. [Blood. 1997]

Recombinant adeno-associated virus-mediated gene transfer into hematopoietic progenitor cells. [Blood. 1994]

Position-independent human beta-globin gene expression mediated by a recombinant adeno-associated virus vector carrying the chicken beta-globin 5' enhancer. [Blood. 1999]

Recombinant adeno-associated virus (rAAV)-mediated expression of a human gamma-globin gene in human progenitor-derived erythroid cells. [Blood. 1994]

Cellular recombination pathways and viral terminal repeat hairpin structures are sufficient for adeno-associated virus integration in vivo and in vitro. [Mol. Cell Biol. 1997]

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